PATENT COOPERATION TREATY

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From the INTERNATIONAL SEARCHING AUTHORITY LOUIS MYERS PCT FISH & RICHARDSON P.C. 225 FRANKLIN STREET BOSTON MA 02110-2804 NOTIFICATION OF TRANSMITTAL OF THE INTERNATIONAL SEARCH REPORT OR THE DECLARATION (PCT Rule 44.1) PC & RICHARDSON **BOSTON OFFICE** Date of Mailing 14 NOV 2000 (day/month/year) Applicant's or agent's file reference FOR FURTHER ACTION 10275=1370+WW)\ See paragraphs 1 and 4 below International application No. International filing date (day/month/year) PCT/US00/25560 18 SEPTEMBER 2000 Applicant GENZYME TRANSGENICS CORPORATION 1. X The applicant is hereby notified that the international search report has been established and is transmitted herewith. Filing of amendments and statement under Article 19: The applicant is entitled, if he so wishes, to amend the claims of the international application (see Rule 46): The time limit for filing such amendments is normally 2 months from the date of transmittal of the international search report; however, for more details, see the notes on the accompanying sheet. Where? Directly to the International Bureau of WIPO 34, chemin des Colombettes 1211 Geneva 20, Switzerland Facsimile No.: (41-22) 740.14.35 For more detailed instructions, see the notes on the accompanying sheet The applicant is hereby notified that no international search report will be established and that Article 17(2)(a) to that effect is transmitted herewith. With regard to the protest against payment of (an) additional fee(s) under Rule 40.2, the applicant is notified that: the protest together with the decision thereon has been transmitted to the International Bureau together with the applicant's request to forward the texts of both the protest and the decision thereon to the designated Offices. no decision has been made yet on the protest; the applicant will be notified as soon as a decision is made. 4. Further action(s): The applicant is reminded of the following: Shortly after 18 months from the priority date, the international application will be published by the International Bureau. If the applicant wishes to avoid or postpone publication, a notice of withdrawal of the international application, or of the priority claim, must reach the International Bureau as provided in rules 90 bis 1 and 90 bis 3, respectively, before the completion of the technical preparations for international publication. Within 19 months from the priority date, a demand for international preliminary examination must be filed if the applicant wishes to postpone the entry into the national phase until 30 months from the priority date (in some Offices even later). Within 20 months from the priority date, the applicant must perform the prescribed acts for entry into the national phase before all designated Offices which have not been elected in the demand or in a later election within 19 months from the priority date or could not be elected because they are not bound by Chapter II. Name and mailing address of the ISA/US Authorized officer Commissioner of Patents and Trademarks JANET M. KERR Form PCT/PS/APS/TOGN 1809 190800

PATENT COOPERATION TREATY

From the INTERNATIONAL SEARCHING AUTHORITY

To: LOUIS MYERS FISH & RICHARDSON P.C. 225 FRANKLIN STREET	PCT			
BOSTON MA 02110-2804	NOTIFICATION OF TRANSMITTAL OF THE INTERNATIONAL SEARCH REPORT OR THE DECLARATION			
·	(PCT Rule 44.1)			
	Date of Mailing (day/month/year) 14 NOV 2000			
Applicant's or agent's file reference 10275-13701W	FOR FURTHER ACTION See paragraphs 1 and 4 below			
International application No. PCT/US00/25560	International filing date (day/month/year) 18 SEPTEMBER 2000			
Applicant GENZYME TRANSGENICS CORPOR	ATION			
The applicant is hereby notified that the international search report has been established and is transmitted herewith. Filing of amendments and statement under Article 19: The applicant is entitled, if he so wishes, to amend the claims of the international application (see Rule 46):				
When? The time limit for filing such amendments is normally 2 months from the date of transmittal of the international search_report; howeverfor_more_details,_see_the_notes_on_the_accompanying sheet.				
Where? Directly to the International Bureau of WIPO 34, chemin des Colombettes 1211 Geneva 20, Switzerland Facsimile No.: (41-22) 740.14.35				
For more detailed instructions, see the notes on the accompanying sheet.				
2. The applicant is hereby notified that no international search report will be established and that the declaration under Article 17(2)(a) to that effect is transmitted herewith.				
3. With regard to the protest against payment of (an) additional fee(s) under Rule 40.2, the applicant is notified that:				
the protest together with the decision thereon applicant's request to forward the texts of bot	has been transmitted to the International Bureau together with the h the protest and the decision thereon to the designated Offices.			
no decision has been made yet on the protest; the applicant-will be notified as soon as a decision is made.				
4. Further action(s): The applicant is reminded of the fol				
the applicant wishes to avoid or postpone publication priority claim, must reach the International Bureau as completion of the technical preparations for internation	•			
Within 19 months from the priority date, a demand for in wishes to postpone the entry into the national phase u	nternational preliminary examination must be filed if the applicant ntil 30 months from the priority date (in some Offices even later).			
Within 20 months from the priority date, the applicant mus all designated Offices which have not been elected in t date or could not be elected because they are not bour	t perform the prescribed acts for entry into the national phase before he demand or in a later election within 19 months from the priority and by Chapter II.			
Name and mailing address of the ISA/US	Authorized officer			
Commissioner of Patents and Trademarks Form PCT 33 PST 1997 PST 19				

INTERNATIONAL SEARCH REPORT

International application No. PCT/US00/25560

A. CLASSIFICATION OF SUBJECT MATTER IPC(7) :C07H 21/04; C07K 1/00, 14/00: C12N 5/00, 15/12. 15/13; C12P 21/00 US CL :Please See Extra Sheet. According to International Patent Classification (IPC) or to both national classification and IPC					
B. FIELDS	S SEARCHED				
Minimum doc	numentation searched (classification system followed	by classification symbols)			
U.S. : 435/69.1, 69.7, 325, 455; 530/350, 387.3; 536/23.1, 23.4; 800/4, 6, 7					
Documentation searched other than minimum documentation to the extent that such documents are included in the fields searched NONE					
Electronic data	a base consulted during the international search (na	me of data base and, where practicable,	search terms used)		
Please See F	Extra Sheet.				
C. DOCU	MENTS CONSIDERED TO BE RELEVANT				
Category*	Citation of document, with indication, where app	propriate, of the relevant passages	Relevant to claim No.		
	NEWTON. D.L. et al. Antitransferrin Receptor Antibody-RNase Fusion Protein Expressed in the Mammary Gland of Transgenic		1,4-8, 10-17		
Y,P 1	Mice. J. Immunol. Methods 1999. Ventire document.	2, 3, 9			
	and the claims.		1, 10, 14-16 		
Y			2-9, 13		
	WO 95/27782 A1 (PPL THERAPEUTICES (SCOTLAND) LTD) 19 October 1995, pages 1-23, see especially pages 17-19.		1, 10, 14-16		
Y			2-9, 13		
			·		
X Further	r documents are listed in the continuation of Box C	See patent family annex.			
• Special categories of cited documents: "A" document defining the general state of the art which is not considered "T" later document published after the international filing date or produce and not in conflict with the application but cited to understart principle or theory underlying the invention		ation but cited to understand the			
1	to be of particular relevance "E" earlier document published on or after the international filing date "X" document of particular relevance; the claimed invention cannot be considered novel or cannot be considered to involve an inventive step				
cited	ment which may throw doubts on priority claim(s) or which is to establish the publication date of another citation or other ial reason (1s specified)	"Y" document of particular relevance; the considered to involve an inventive	e claimed invention cannot be step when the document is		
"P" docur	ment referring to an oral disclosure, use, exhibition or other means	combined with one or more other suc being obvious to a person skilled in t	h documents, such combination he art		
·	ctual completion of the international search	"&" document member of the same patent Date of mailing of the international sea			
20 OCTOBER 2000 14 NOV 2000 17		2000 /7			
Name and mailing address of the ISA/US Hoston ble to a ddress of the ISA/US Box PCT Authorized officer Telephone No. (703) 308-0196 Authorized officer Telephone No.					

INTERNATIONAL SEARCH REPORT

International application No. PCT/US00/25560

Category*	Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No
X,P	US 5,959,171 A (HYTTINEN et al.) 28 September 1999, col. 6 to col. 10.	1, 10-12, 14-17
Y,P		2-9, 13
Y	US 5,840,840 (RYBAK et al.) 24 November 1998, col. 7 to col. 18.	6-8
Y	US 5,206,161 (DRAYNA et al.) 27 April 1993, col. 34 to col. 38.	9
7	US 5,948,668 A (HARTMAN et al.) 07 September 1999, col. 2 to col. 10.	9
7	WO 98/18809 A1 (CELL GENESYS, INC.) 07 May 1998, pages 1-62, see especially pages 8, 12-16, and 21.	1, 3, 5
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INTERNATIONAL SEARCH REPORT

International application No. PCT/US00/25560

A. CLA	SSIFICATION	OF SUBJEC	CT MATTER:
US CL	:		

435/69.1, 69.7, 325, 455; 530/350, 387.3; 536/23.1, 23.4; 800/4, 6, 7

B. FIELDS SEARCHED

Electronic data bases consulted (Name of data base and where practicable terms used):

WEST, MEDLINE, EMBASE, BIOSIS, INPADOC, CAPLUS

search terms: transgenic, fusion protein, chimeric, ig, igg, tag-72, carcinoembryonic antigen, rnase, rnase a, carboxypeptidase, angiogenin, transferrin receptor, milk

Form PCT/ISA/210 (extra sheet) (July 1998)*

NOTES TO FORM PCT/ISA/220

These Notes are intended to give the basic instructions concerning the Gling of emendments under Article 19. The Notes are based on the requirements of the Patent Cooperation Treaty and of the Reguladess and the Administrative frustructions under that Treaty. In case of discrepancy between these Notes and those requirements, the latter are applicable. For more detailed information, see also the PCT Applicant's Quide, a publication of WIPO.

In these Notes, "Article", "Rule" and "Section" refer to the provisions of the PCT, the PCT Regulations and the PCT Administrative Instructions, respectively.

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INSTRUCTIONS CONCERNING AMENDMENTS UNDER ARTICLE 19

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The applicant has, other having received the innecessional search report, one opportunity to amend the claims of the intermedicant application. It should however be complexized that, since all parts of the intermedical application (claims, description and derivings) may be amended during the intermedical pediamenty extended the precedure, there is versely no need to file amendments of the claims under Asticle 19 except where, e.g. the applicant weak the latter to be published for the purposes of provisional protection or has another reason for amending the claims before interestional publication. Purchamore, it should be complexized that provisional protection is available in some States only.

What parts of the international application may be assended?

The claims only.

The description and the derivings may only be amended during inticastional preliminary examination under Chapter II.

When? Within 2 meeths from the date of transmitted of the international courts report or 16 meeths from the priority date, whichever these limit expires later. It should be noted, however, that the amendments will be equivious or bring been received on time If they are received by the international Buress offer the expiration of the expiration fine limit but before the completion of the technical proposalous for international publication (finis 46.1).

Where not to the the emendments?

The amendments may only be filed with the International Bureau and por with the seceiving Office or the International Searching Authority (Rule 44.2).

When a demand for international purlimic sy examination has been a filed, see below.

Eow ? Either by cancelling one or more entire claims, by adding one or more new claims or by smeading the test of one or more of the claims as filed.

A replacement short must be submitted for each short of the claims which, on account of an amendment or amendments, differs from the short originally filed.

All the claims appearing on a replacement short must be numbered in Arabic numerals. Where a claim is cancelled, no renumbering of the other claims is required. In all cases where claims are renumbered, they must be renumbered consecutively (Administrative Instructions, Section 205(b)).

What documents must/may accompany the amendments?

Letter (Section 205(b)):

The emendments must be submitted with a letter.

The letter will not be published with the international application and the emended claims. It should not be confounded with the "Statement under Article 19(1)" (see below, under "Statement under Article 19(1)").

The letter must indicate the differences between the claims as filed and the claims as amended. It must, in particular, indicate, in connection with each claim appearing in the international application (it being understood that identical indications concerning several claims may be grouped), whether

- (i) the claim is unchanged;
- (ii) the claim is cancelled;
- (iii) the claim is new:
- (iv) the claim replaces one or more claims as filed;
- (v) the claim is the result of the division of a claim as filed.